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## LOGINID:ssspta1604dxj

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TERMINAL (ENTER 1, 2, 3, OR ?):2

* * *	* *	* *	* *	* Welcome to STN International * * * * * * * * *
NEWS	1			Web Page for STN Seminar Schedule - N. America
NEWS	2	NOV	21	CAS patent coverage to include exemplified prophetic substances identified in English-, French-, German-, and Japanese-language basic patents from 2004-present
NEWS	3	NOV	26	MARPAT enhanced with FSORT command
NEWS	4	NOV	26	CHEMSAFE now available on STN Easy
NEWS	5	NOV	26	Two new SET commands increase convenience of STN searching
NEWS	6	DEC	0.1	ChemPort single article sales feature unavailable
NEWS	7	DEC		GBFULL now offers single source for full-text
				coverage of complete UK patent families
NEWS	8	DEC	17	Fifty-one pharmaceutical ingredients added to PS
NEWS	9	JAN	06	The retention policy for unread STNmail messages
				will change in 2009 for STN-Columbus and STN-Tokyo
NEWS	10	JAN	07	WPIDS, WPINDEX, and WPIX enhanced Japanese Patent
				Classification Data
NEWS	11	FEB	02	Simultaneous left and right truncation (SLART) added for CERAB, COMPUAB, ELCOM, and SOLIDSTATE
NEWS	1.2	FEB	02	GENBANK enhanced with SET PLURALS and SET SPELLING
NEWS	13	FEB	06	Patent sequence location (PSL) data added to USGENE
NEWS	14	FEB	10	COMPENDEX reloaded and enhanced
NEWS	15	FEB	11	WTEXTILES reloaded and enhanced
NEWS	16	FEB	19	New patent-examiner citations in 300,000 CA/CAplus
				patent records provide insights into related prior art
NEWS	17	FEB	19	Increase the precision of your patent queries use terms from the IPC Thesaurus, Version 2009.01
NEWS	18	FEB	23	Several formats for image display and print options discontinued in USPATFULL and USPAT2
NEWS	19	FEB	23	MEDLINE now offers more precise author group fields
				and 2009 MeSH terms
NEWS	20	FEB	23	TOXCENTER updates mirror those of MEDLINE - more
NEWS	21	FEB	22	precise author group fields and 2009 MeSH terms Three million new patent records blast AEROSPACE into
NENS	21	FED	23	STN patent clusters
NEWS	22	FEB	25	USGENE enhanced with patent family and legal status
				display data from INPADOCDB
NEWS	23	MAR	06	INPADOCDB and INPAFAMDB enhanced with new display formats
NEWS	EXP	RESS		E 27 08 CURRENT WINDOWS VERSION IS V8.3,
			AND	CURRENT DISCOVER FILE IS DATED 23 JUNE 2008.

NEWS HOURS STN Operating Hours Plus Help Desk Availability

NEWS LOGIN Welcome Banner and News Items

NEWS IPC8 For general information regarding STN implementation of IPC 8

Enter NEWS followed by the item number or name to see news on that specific topic.

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SINCE FILE

ENTRY

0.22

TOTAL

0.22

SESSION

FILE 'HOME' ENTERED AT 15:26:56 ON 09 MAR 2009

=> FIL REGISTRY

COST IN U.S. DOLLARS

FULL ESTIMATED COST

FILE 'REGISTRY' ENTERED AT 15:27:10 ON 09 MAR 2009
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STRUCTURE FILE UPDATES: 8 MAR 2009 HIGHEST RN 1117698-24-4
DICTIONARY FILE UPDATES: 8 MAR 2009 HIGHEST RN 1117698-24-4

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TSCA INFORMATION NOW CURRENT THROUGH January 9, 2009.

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http://www.cas.org/support/stngen/stndoc/properties.html

=> s strontium ranelate 77648 STRONTIUM

2 RANELATE

1 STRONTIUM RANELATE

(STRONTIUM(W)RANELATE)

=> s ranelate L2 2 RANELATE

=> d 12 1-2

L2 ANSWER 1 OF 2 REGISTRY COPYRIGHT 2009 ACS on STN

RN 135459-87-9 REGISTRY

ED Entered STN: 09 Aug 1991

```
10533787
     3-Thiopheneacetic acid, 5-[bis(carboxymethyl)amino]-2-carboxy-4-cyano-,
     strontium salt (1:2) (CA INDEX NAME)
OTHER NAMES:
CN
    Distrontium renelate
     Protelos
CN
     Protos
CN
    S 12911
CN
     S 12911-2
CN
     Strontium ranelate
     C12 H10 N2 O8 S . 2 Sr
MF
SR
     CA
LC
     STN Files: ADISINSIGHT, AGRICOLA, ANABSTR, BIOSIS, CA, CAPLUS, CASREACT,
      CHEMCATS, CIN, EMBASE, IMSDRUGNEWS, IMSPATENTS, IMSPRODUCT, IMSRESEARCH,
       IPA, MRCK*, PATDPASPC, PHAR, PROMT, PROUSDDR, SYNTHLINE, TOXCENTER,
       USPAT2, USPATFULL
         (*File contains numerically searchable property data)
     Other Sources:
                    WHO
CRN (135459-90-4)
     HO2C-CH2
HO2C-CH2-N-
                    CO2H
                   сно-соон
```

●2 Sr

173 REFERENCES IN FILE CA (1907 TO DATE)

1 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
173 REFERENCES IN FILE CAPLUS (1907 TO DATE)

- L2 ANSWER 2 OF 2 REGISTRY COPYRIGHT 2009 ACS on STN
- RN 58194-26-6 REGISTRY
- ED Entered STN: 16 Nov 1984
- CN 3-Thiopheneacetic acid, 5-[bis(2-ethoxy-2-oxoethyl)amino]-4-cyano-2-(ethoxycarbonyl)-, ethyl ester (CA INDEX NAME)

OTHER NAMES:

- CN <u>Tetraethyl</u> ranelate MF C20 H26 N2 O8 S
- LC STN Files: BEILSTEIN\*, CA, CAPLUS, CASREACT, CHEMCATS, TOXCENTER, USPATFULL

(\*File contains numerically searchable property data)

\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

7 REFERENCES IN FILE CA (1907 TO DATE)

7 REFERENCES IN FILE CAPLUS (1907 TO DATE)

SINCE FILE

ENTRY

21.11

TOTAL

21.33

SESSION

=> file medicine FILE 'DRUGMONOG' ACCESS NOT AUTHORIZED

COST IN U.S. DOLLARS

FULL ESTIMATED COST

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FILE 'USPATFULL' ENTERED AT 15:28:07 ON 09 MAR 2009
CA INDEXING COPYRIGHT (C) 2009 AMERICAN CHEMICAL SOCIETY (ACS)

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FILE 'USPATOLD' ENTERED AT 15:28:07 ON 09 MAR 2009
CA INDEXING COPYRIGHT (C) 2009 AMERICAN CHEMICAL SOCIETY (ACS)
FILE 'USPAT2' ENTERED AT 15:28:07 ON 09 MAR 2009
CA INDEXING COPYRIGHT (C) 2009 AMERICAN CHEMICAL SOCIETY (ACS)
=> s 12 or ranelate
         2655 L2 OR RANELATE
1.3
=> s protelos or protos
         1034 PROTELOS OR PROTOS
=> s 13 or 14
L5
          3305 L3 OR L4
=> s pain or ?itis
LEFT TRUNCATION IGNORED FOR FILE 'ADISINSIGHT'
LEFT TRUNCATION IGNORED FOR FILE 'ADISNEWS'
   5 FILES SEARCHED...
LEFT TRUNCATION IGNORED FOR FILE 'DDFB'
LEFT TRUNCATION IGNORED FOR FILE 'DGENE'
LEFT TRUNCATION IGNORED FOR FILE 'DRUGB'
LEFT TRUNCATION IGNORED FOR FILE 'DRUGMONOG2'
LEFT TRUNCATION IGNORED FOR FILE 'DRUGU'
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LEFT TRUNCATION IGNORED FOR FILE 'IPA'
LEFT TRUNCATION IGNORED FOR FILE 'LIFESCI'
LEFT TRUNCATION IGNORED FOR FILE 'NLDB'
LEFT TRUNCATION IGNORED FOR FILE 'NUTRACEUT'
 26 FILES SEARCHED...
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LEFT TRUNCATION IGNORED FOR FILE 'PHARMAML'
 31 FILES SEARCHED...
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LEFT TRUNCATION IGNORED FOR FILE 'USPATOLD'
LEFT TRUNCATION IGNORED FOR FILE 'USPATOLD'
LEFT TRUNCATION IGNORED FOR FILE 'USPAT2'
LEFT TRUNCATION IGNORED FOR FILE 'USPAT2'
LEFT TRUNCATION IGNORED FOR FILE 'USPAT2'
       8213771 PAIN OR ?ITIS
Left truncation is not valid in the specified search field in the
specified file. The term has been searched without left truncation.
Examples: '?TERPEN?' would be searched as 'TERPEN?' and '?FLAVONOID'
would be searched as 'FLAVONOID.'
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If you are searching in a field that uses implied proximity, and you used a truncation symbol after a punctuation mark, the system may interpret the truncation symbol as being at the beginning of a term. Implied proximity is used in search fields indexed as single words, for example, the Basic Index.

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L7 519 L5 AND L6
=> s 17 and pd<2003
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=> s 15 and 16

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5 FILES SEARCHED...
'2003' NOT A VALID FIELD CODE
'2003' NOT A VALID FIELD CODE
'2003' NOT A VALID FIELD CODE
  15 FILES SEARCHED...
'2003' NOT A VALID FIELD CODE
 22 FILES SEARCHED...
'2003' NOT A VALID FIELD CODE
 28 FILES SEARCHED...
'2003' NOT A VALID FIELD CODE
 31 FILES SEARCHED...
          17 L7 AND PD<2003
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ENTER L# LIST OR (END):18
DUPLICATE IS NOT AVAILABLE IN 'ADISINSIGHT, ADISNEWS, DGENE, DRUGMONOG2,
IMSPRODUCT, KOSMET, NUTRACEUT, PCTGEN, PHARMAML, USGENE'.
ANSWERS FROM THESE FILES WILL BE CONSIDERED UNIOUE
PROCESSING COMPLETED FOR L8
1.9
            13 DUP REM L8 (4 DUPLICATES REMOVED)
=> d 19 1-13 ibib, kwic
L9 ANSWER 1 OF 13 USPATFULL on STN
ACCESSION NUMBER:
                       2002:343531 USPATFULL
TITLE:
                       Soluble lymphotoxin beta receptor and anti-lymphotoxin
                       receptor and ligand antibodies as therapeutic agents
                       for treatment
INVENTOR(S):
                       Browning, Jeffrey L., Brookline, MA, UNITED STATES
                       Hochman, Paula S., Newton, MA, UNITED STATES
                       Rennert, Paul D., Millis, MA, UNITED STATES
                       MacKay, Fabienne, Vaucluse, AUSTRALIA
                            NUMBER
                                        KIND DATE
                       -----
                       US 20020197254 A1 20021226

US 7309492 B2 20071218

US 2001-3211 A1 20011031 (10)
PATENT INFORMATION:
APPLICATION INFO.:
RELATED APPLN. INFO.: Continuation of Ser. No. US 1999-299139, filed on 23
                       Apr 1999, PENDING
                             NUMBER DATE
PRIORITY INFORMATION: WO 1997-US19436 19971024
                      US 1996-29060P
                                        19961025 (60)
DOCUMENT TYPE:
                      Utility
FILE SEGMENT:
                       APPLICATION
LEGAL REPRESENTATIVE: Niki D. Cox, Esq., BIOGEN, INC., 14 Cambridge Center,
                       Cambridge, MA, 02142
NUMBER OF CLAIMS:
                       50
EXEMPLARY CLAIM:
                       1
NUMBER OF DRAWINGS:
                      10 Drawing Page(s)
LINE COUNT:
                       2115
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       . . . which in turn activates mast cells to produce acute
       inflammatory reactions such as those which lead to eczema, asthma and
```

. . . immune responses are associated with a number of organ-specific and systemic autoimmune conditions such as Systemic Lupus Erythematosus,

SUMM

Wegener's Granulomatosis, Polyarteritis Nodosa (PAN), Rapidly Progressive Crescentic Glomerulonephritis and Idiopathic Thrombocytopenia Purpura, as well as chronic inflammatory diseases such as the Graves' and Chagas' disease. Humoral immune responses, , .

. . . caused by molecular mimicry. For example, the immune reaction DETD to the lyme disease infectious agent Borrelia burgdorferi leads to an arthritis-like disease presumably because saome antigenic epitope on this bacterium resembles a normal joint component. Removal of the FDC-retained lyme bacterium antigen may ameliorate lyme disease induced arthritis. Such therapy would also be relevant to other cases of mimicry associated with infectious agents.

DETD . . Miller et al., J. Exp. Med., 178, pp. 211-222 (1993)). Purified human IgG1 used as a control was purchased from Protos Immunoresearch (San Francisco, Calif.). MR1, anti-mouse CD40 ligand antibody, was purchased from Pharmingen (San Diego, Calif.).

DETD . . include: Myasthenia Gravis, autoimmune hemolytic anemia, Chagas' disease, Grave's disease, idiopathic thrombocytopenia purpura (ITP) Systemic Lupus Erythematosus (SLE), Wegener's Granulomatosis, Poly-arteritis Nodosa and Rapidly Progressive Crescentic Glomerulonephritis. (From Benjamini, et al. Immunology, A Short Course, (Wiley-Liss, New York 3d ed. (1996)) Although the etiology of SLE is. . . in joint synovial spaces. These complexes activate the complement cascade and attract granulocytes. The subsequent inflammatory reaction is characterized as glomerulonephritis, with resulting damage to the kidneys leading to proteinuria and hematuria.

DETD [0174] Lupus nephritis has been studied in murine models for decades. Recently, the therapeutic efficacy of a reagent specific for the murine CD40.

DETD . . of activation, and damage from the release of lytic enzymes from their granules results in the destruction of cells. Rheumatic arthritis is thought to result from a type III hypersensitivity reaction mediated by immune complexes of antigen (in this case rheumatoid. .

DETD . . . reagent which inhibits antibody responses to ameliorate a pathologic immunological response is supported in the recent study of mouse lupus nephritis. In the latter study, administration of an antibody that blocks the CD40/CD40L pathway was shown inhibit the acceleration of lupus nephritis produced upon transfer of cells which induce the production of pathogenic antibodies in vivo, and have a sustained beneficial effect.

ANSWER 2 OF 13 USPATFULL on STN ACCESSION NUMBER:

2002:272935 USPATFULL

TITLE: Novel differentiation inducing process of embryonic stem cell to ectodermal cell and its use

INVENTOR(S): Sasai, Yoshiki, Kyoto, JAPAN

Nishikawa, Shin-Ichi, Kyoto, JAPAN

	NUMBER	KIND	DATE		
PATENT INFORMATION:	US 20020151056	A1	20021017		<
APPLICATION INFO.:	US 2001-855587	Al	20010516	(9)	

NUMBER DATE PRIORITY INFORMATION: JP 2000-144059 20000516 JP 2000-290819 20000925 US 2000-257049P 20001220 (60) DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

Jagoe

```
LEGAL REPRESENTATIVE: FITZPATRICK CELLA HARPER & SCINTO, 30 ROCKEFELLER
                      PLAZA, NEW YORK, NY, 10112
NUMBER OF CLAIMS:
                      71
EXEMPLARY CLAIM:
                      1
                     10 Drawing Page(s)
NUMBER OF DRAWINGS:
LINE COUNT:
                      4056
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
           . so far been established in rat (P. M. Iannaccone et al., Dev.
      Biol., 163, 288 (1994)), in domestic fowl (B. Pain et al.,
      Development, 122, 2339 (1996); U.S. Pat. No. 5,340,740; U.S. Pat. No.
      5,656,479)), in pig (M. B. Wheeler, Reprod.. . .
       . . serotonergic neuron marker serotinin (manufactured by Dia
      Sorin) or an antibody against a noradrenaline neuron marker dopamine
      β-hydroxylase (manufactured by PROTOS Biotech).
L9 ANSWER 3 OF 13 USPATFULL on STN
ACCESSION NUMBER:
                    2002:243628 USPATFULL
TITLE:
                      Novel purinse
INVENTOR(S):
                      Metcalf, Chester A., III, Boston, MA, UNITED STATES
                      Weigele, Manfred, Cambridge, MA, UNITED STATES
                      Sawyer, Tomi K., Southborough, MA, UNITED STATES
                      Bohacek, Regine, Boston, MA, UNITED STATES
                      Shakespeare, William C., Framingham, MA, UNITED STATES
                      Sundaramoorthi, Rajeswari, Watertown, MA, UNITED STATES
                      Wang, Yihan, Newton, MA, UNITED STATES
                      Dalgarno, David C., Brookline, MA, UNITED STATES
                           NUMBER
                                    KIND DATE
                      _____
PATENT INFORMATION:
                      US 20020132819 A1 20020919
                                                               <---
APPLICATION INFO.:
                      US 2000-740653
                                       A1 20001218 (9)
                            NUMBER
                                        DATE
                      -----
PRIORITY INFORMATION:
                      US 1999-172510P 19991217 (60)
                      US 1999-172161P
                                        19991217 (60)
                      US 2000-240788P 20001016 (60)
DOCUMENT TYPE:
                      Utility
FILE SEGMENT:
                      APPLICATION
LEGAL REPRESENTATIVE: Karoline Shair, Ph.D., Choate, Hall & Stewart, 53 State
                      Street, Exchange Place, Boston, MA, 02109
NUMBER OF CLAIMS:
                      195
EXEMPLARY CLAIM:
                     1
                      4673
LINE COUNT:
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
      . . . limited to, Paget's Disease, primary and secondary
      hyperparathyroidism, humoral hypercalcemia of malignancy, various
      cancers where resorption is increased, and rheumatoid arthritis
SHMM
      . . fast increase in bone mineral content by promoting osteoblast
      activity. Such examples include peptides from the parathyroid hormone
      family, strontium ranelate, and growth hormone and
      insulin-like growth response (see, for example, Reginster et al.
      "Promising New Agents in Osteoporosis," Drugs R. . .
   ANSWER 4 OF 13 USPATFULL on STN
ACCESSION NUMBER: 2002:192090 USPATFULL
```

Novel heterocycles

TITLE:

INVENTOR(S):

Weigele, Manfred, Cambridge, MA, UNITED STATES Luke, George P., Clinton, CT, UNITED STATES Sawyer, Tomi K., Southborough, MA, UNITED STATES Bohacek, Regine, Boston, MA, UNITED STATES Shakespeare, William C., Framingham, MA, UNITED STATES Sundaramoorthi, Rajeswari, Watertown, MA, UNITED STATES Wang, Yihan, Newton, MA, UNITED STATES Dalgarno, David C., Brookline, MA, UNITED STATES Metcalf, Chester A., III, Boston, MA, UNITED STATES

Vu, Chi B., Arlington, MA, UNITED STATES Kawahata, Noriyuki H., Medford, MA, UNITED STATES KIND DATE

-----PATENT INFORMATION: US 20020103161 A1 20020801 <--US 2000-740267 A1 20001218 (9) APPLICATION INFO.: NUMBER DATE US 1999-172510P 19991217 (60) US 1999-172161P 19991217 (60) US 2000-240788P 20001016 (60) PRIORITY INFORMATION:

NUMBER

DOCUMENT TYPE: Utility

FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: Karoline K.M. Shair, Ph.D., Choate, Hall & Stewart, 53 State Street, Exchange Place, Boston, MA, 02109

NUMBER OF CLAIMS: 111 EXEMPLARY CLAIM: 1

LINE COUNT: 4552

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

. . . limited to, Paget's Disease, primary and secondary hyperparathyroidism, humoral hypercalcemia of malignancy, various cancers where resorption is increased, and rheumatoid arthritis

. fast increase in bone mineral content by promoting osteoblast SUMM activity. Such examples include peptides from the parathyroid hormone family, strontium ranelate, and growth hormone and insulin-like growth response (see, for example, Reginster et al. "Promising New Agents in Osteoporosis," Drugs R. . SUMM . . . sterile isotonic aqueous buffer. Where necessary, the

composition may also include a solubilizing agent and a local anesthetic to ease pain at the side of the injection. Generally, the ingredients are supplied either separately or mixed together in unit dosage form, . .

L9 ANSWER 5 OF 13 USPATFULL on STN

ACCESSION NUMBER: 2002:191593 USPATFULL

TITLE: Human monoclonal antibody against a costimulatory

signal transduction molecule AILIM and pharmaceutical

use thereof

INVENTOR(S): Tsuji, Takashi, Nagareyama-shi, JAPAN

Tezuka, Katsunari, Yokohama-shi, JAPAN Hori, Nobuaki, Yokohama-shi, JAPAN

	NUMBER	KIND	DATE	
PATENT INFORMATION:	US 20020102658	A1	20020801	
	US 6803039	B2	20041012	
APPLICATION INFO.:	US 2001-859053	A1	20010516	(

NUMBER DATE JP 2000-147116 20000518 PRIORITY INFORMATION: JP 2001-99508 20010330 DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION JANIS K. FRASER, PH.D., J.D., Fish & Richardson P.C., LEGAL REPRESENTATIVE: 225 Franklin Street, Boston, MA, 02110-2804 NUMBER OF CLAIMS: 108 EXEMPLARY CLAIM: NUMBER OF DRAWINGS: 78 Drawing Page(s) LINE COUNT: 6932 CAS INDEXING IS AVAILABLE FOR THIS PATENT. . . . of pharmaceutical compositions according to this invention enables suppression, prevention and/or treatment of, for example, various disorders (for example, rheumatoid arthritis, multiple sclerosis, autoimmune thyroiditis, allergic contact-type dermatitis, chronic inflammatory dermatosis such as lichen planus, systemic lupus erythematosus, insulin-dependent diabetes mellitus, psoriasis, etc.) classified into autoimmune or allergic disorders (particularly autoimmune disease and delayed allergy caused by cellular immunity); arthropathia (for example, rheumatoid arthritis (RA) and osteoarthritis (OA)), inflammation (e.g., hepatitis); graft versus host reaction (GVH reaction); graft versus host disease (GVHD); immune rejection accompanying transplantation (homoplasty or heteroplasty) of a. . . of cytokines); and disorders possibly caused by the abnormal intestinal immunity (specifically inflammatory intestinal disorders (particularly clone disease and ulcerative colitis) and alimentary allergy). SUMM [0030] The pharmaceutical composition of the present invention can be applied to inflammatory disease for example, inflammation accompanying various arthritis (for example, rheumatoid arthritis , osteoarthritis), pneumonia, hepatitis (including viral hepatitis), inflammation accompanying infectious inflammatory bowel diseases, intestinal enteritis, nephritis (inflammation accompanying glomerular nephritis, nephrofibrosis), gastritis, angiitis, pancreatitis, peritonitis, bronchitis, myocarditis, cerebritis, inflammation in postischemic reperfusion injury (myocardial ischemic reperfusion in urv), inflammation attributed to immune rejection after transplantation of tissue and organ, burn, various skin inflammation (psoriasis, allergic contact-type dermatitis, lichen planus which is chronic inflammatory skin disease), inflammation in multiple organ failure, inflammation after operation of PTCA or PTCR, and inflammation accompanying arteriosclerosis, and autoimmune thyroiditis. . . . an active ingredient, it is possible to inhibit or treat and prevent, for example, a variety of diseases (e.g., rheumatoid DETD arthritis, multiple sclerosis, autoimmune thyroiditis, allergic contact dermatitis, lichen planus as a chronic inflammatory skin disease, systemic lupus erythematosus, insulin dependent diabetes mellitus and psoriasis, etc.) classified into autoimmune diseases or allergic diseases (particularly, autoimmune diseases and delayed allergies by cellular immunity); arthropathies (e.g., rheumatoid arthritis (RA), osteoarthritis (OA)), inflammation (e.g., hepatitis); graft versus host reaction (GVH reaction); graft versus host disease (graft versus host

disease; GVHD); immunorejection associated with transplantation (allogenic. . . and diseases that are potentially caused by abnormality in gut immunity (specifically, inflammatory bowel disease (particularly, Crohn's disease and ulcerative colitis); and alimentary allergy, etc.

DETD . . . some inflammations for which various steroidal drugs are used as anti-inflammatory drugs, for example, inflammation associated with various arthritides (rheumatoid arthritis,

osteoarthritis, etc.), pneumonia, hepatitis (including viral hepatitis), inflammation associated with infectious diseases, inflammatory bowel disease, enteritis, nephritis (glomerular nephritis, inflammation

associated with kidney fibrosis, gastritis, vasculitis

, pancreatitis, peritonitis, bronchitis,

myocarditis, encephalitis, inflammation associated with ischemia-reperfusion injury (myocaridial ischemia-reperfusion injury, etc.), inflammation associated with immunorejection after transplantation of tissues or organs, scald, various skin inflammations (psoriasis, allergic contact dermatitis, lichen planus as a chronic inflammatory skin disease), inflammation associated with multiple organ failure, inflammation after operation of PTCA or PTCR, and inflammation associated with atherosclerosis, autoimmune

thyroiditis, etc. DETD [1008] Biotin-labeled anti-human IgG antibody (Protos);

[1041] Subsequently, peroxidase-conjugated goat anti-human IgG/k antibody was added to each well (4,000 times diluted, 100 µl/well, Protos), and the plate was incubated at room temperature for 1 hour

L9 ANSWER 6 OF 13 USPATFULL on STN

2002:133863 USPATFULL ACCESSION NUMBER:

DETD

TITLE: Purine derivatives

INVENTOR(S): Weigele, Manfred, Cambridge, MA, UNITED STATES Sawyer, Tomi K., Southborough, MA, UNITED STATES Bohacek, Regine, Boston, MA, UNITED STATES

> Shakespeare, William C., Framingham, MA, UNITED STATES Sundaramoorthi, Rajeswari, Watertown, MA, UNITED STATES Wang, Yihan, Newton, MA, UNITED STATES

Dalgarno, David C., Brookline, MA, UNITED STATES Metcalf, Chester A., III, Boston, MA, UNITED STATES

NUMBER KIND DATE US 20020068721 A1 20020606 PATENT INFORMATION: US 7115589 B2 20061003 US 2000-740393 A1 20001218 (9) APPLICATION INFO.:

Continuation-in-part of Ser. No. US 2000-740267, filed RELATED APPLN. INFO.: on 18 Dec 2000, PENDING Continuation-in-part of Ser. No. US 2000-740653, filed on 18 Dec 2000, PENDING

DATE NUMBER US 2000-240788P 20001016 (60) PRIORITY INFORMATION: US 1999-172161P 19991217 (60) US 1999-172510P 19991217 (60) DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: David L. Berstein, ARIAD Pharmaceuticals, Inc., 26 Landsdowne Street, Cambridge, MA, 02139-4234

NUMBER OF CLAIMS:

Jagoe

```
EXEMPLARY CLAIM:
LIME COUNT:
                        3811
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
         . . limited to, Paget's Disease, primary and secondary
       hyperparathyroidism, humoral hypercalcemia of malignancy, various
       cancers where resorption is increased, and rheumatoid arthritis
SHIMM
              fast increase in bone mineral content by promoting osteoblast
       activity. Such examples include peptides from the parathyroid hormone
       family, strontium ranelate, and growth hormone and
       insulin-like growth response (see, for example, "Promising New Agents in
       Osteoporosis", Reginster et al. Drugs R.
     ANSWER 7 OF 13 EMBASE COPYRIGHT (c) 2009 Elsevier B.V. All rights
     reserved on STN
ACCESSION NUMBER:
                   2002177679 EMBASE
TITLE:
                    Strontium ranelate: Dose-dependent effects in
                    established postmenopausal vertebral osteoporosis - A
                    2-year randomized placebo controlled trial.
AUTHOR:
                    Meunier, Pierre J., Dr. (correspondence); Slosman, D.O.;
                    Delmas, P.D.; Sebert, J.L.; Brandi, M.L.; Albanese, C.;
                    Lorenc, R.; Pors-Nielsen, S.; De Vernejoul, M.C.; Roces,
                    A.; Reginster, J.Y.
                    Hopital Edouard Herriot, 69437 Lyon Cedex 03, France.
CORPORATE SOURCE:
                    Meunier@lyon151.inserm.fr
SOURCE:
                    Journal of Clinical Endocrinology and Metabolism, (
                    2002) Vol. 87, No. 5, pp. 2060-2066.
                    Refs: 24
                    ISSN: 0021-972X CODEN: JCEMAZ
COUNTRY:
                    United States
DOCUMENT TYPE:
                    Journal; Article
FILE SEGMENT:
                    017
                           Public Health, Social Medicine and Epidemiology
                    003
                           Endocrinology
                    033
                            Orthopedic Surgery
                            Drug Literature Index
                    0.38
                            Adverse Reactions Titles
LANGUAGE:
                    English
SUMMARY LANGUAGE: English
ENTRY DATE:
                    Entered STN: 6 Jun 2002
                    Last Undated on STN: 6 Jun 2002
     Strontium ranelate: Dose-dependent effects in established
     postmenopausal vertebral osteoporosis - A 2-year randomized placebo
     controlled trial.
     Journal of Clinical Endocrinology and Metabolism, (2002) Vol.
     87, No. 5, pp. 2060-2066.
     Refs: 24
     ISSN: 0021-972X CODEN: JCEMAZ
     The aim of the strontium ranelate (SR) for treatment of
     osteoporosis (STRATOS) trial was to investigate the efficacy and safety of
     different doses of SR, a.
     Medical Descriptors:
       abdominal pain: SI, side effect
     adult
     aged
     alkaline phosphatase blood level
     arthralgia: SI, side effect
     article
     asthenia: SI, side effect
     backache: SI, side effect
```

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bone density
     bone metabolism
    bone mineral
      bronchitis: SI, side effect
     clinical trial
     confidence interval
     controlled study
     coughing: SI, side effect
    dose response
    double blind procedure
    drug efficacy
    drug safety
    drug tolerability
     dual energy X ray absorptiometry
     female
     gastrointestinal symptom: SI, side effect
    human
    hypertension: SI, side effect
     lumbar spine
    major clinical study
     multicenter study
    myalgia: SI, side effect
    neuralgia: SI, side effect
    osteolysis
    outcomes research
      pharyngitis: SI, side effect
     *postmenopause osteoporosis: DT, drug therapy
    priority journal
     randomized controlled trial
       rhinitis: SI, side effect
     vertebra fracture
    vertebra malformation
     vertigo: SI, side effect
     alkaline phosphatase: EC, endogenous compound
     amino terminal telopeptide: EC, endogenous compound
     peptide: EC, endogenous compound
     *strontium: AE, adverse drug reaction
     *strontium: CT, clinical trial
     *strontium: DO, drug dose
     *strontium: DT, drug therapy
       *strontium ranelate: AE, adverse drug reaction
       *strontium ranelate: CT, clinical trial
       *strontium ranelate: DO, drug dose
       *strontium ranelate: DT, drug therapy
    unclassified drug
     (alkaline phosphatase) 9001-78-9; (strontium ranelate)
     135459-87-9; (strontium) 7440-24-6
    ANSWER 8 OF 13 EMBASE COPYRIGHT (c) 2009 Elsevier B.V. All rights
     reserved on STN
                   2002221691 EMBASE
ACCESSION NUMBER:
TITLE:
                    Treatment of postmenopausal osteoporosis.
AUTHOR:
                    Delmas, Pierre D., Dr. (correspondence)
CORPORATE SOURCE:
                  Claude Bernard University of Lyon, France. delmas@lyon151.i
                    nserm.fr
                    Delmas, Pierre D., Dr. (correspondence)
CORPORATE SOURCE:
                    INSERM Research Unit 403, Lyon, France, delmas@lyon151.inse
AUTHOR:
                    Delmas, Pierre D., Dr. (correspondence)
CORPORATE SOURCE: Hopital e Herriot, Pavillon F, 69437 Lyon Cedex 03, France.
```

PМ

```
delmas@lyon151.inserm.fr
AUTHOR:
                    Delmas, Pierre D., Dr. (correspondence)
CORPORATE SOURCE:
                   Hopital E Herriot, Pavillon F, 69437 Lyon Cedex 03, France.
                   delmas@lyon151.inserm.fr
                    Lancet, (8 Jun 2002) Vol. 359, No. 9322, pp.
SOURCE:
                    2018-2026
                    Refs: 111
                    ISSN: 0140-6736 CODEN: LANCAO
COUNTRY:
                   United Kingdom
                   Journal; Article
DOCUMENT TYPE:
FILE SEGMENT:
                   010
                           Obstetrics and Gynecology
                   030
                           Clinical and Experimental Pharmacology
                   033
                           Orthopedic Surgery
                   037
                          Drug Literature Index
                   038
                           Adverse Reactions Titles
LANGUAGE:
                   English
SUMMARY LANGUAGE:
                   English
ENTRY DATE:
                   Entered STN: 11 Jul 2002
                    Last Updated on STN: 11 Jul 2002
     Lancet, (8 Jun 2002) Vol. 359, No. 9322, pp. 2018-2026.
     Refs: 111
     ISSN: 0140-6736 CODEN: LANCAO
    Medical Descriptors:
    age
    article
    bone density
    bone mineral
    calcium intake
    clinical trial
    cognitive defect: SI, side effect
    diarrhea: SI, side effect
    diet
    drug efficacy
    drug induced disease: SI, side effect
    elderly care
      esophagitis: SI, side effect
     exercise
     falling
     flushing
     *fracture: DT, drug therapy
     *fracture: PC, prevention
     gastrointestinal disease: SI, side effect
     *hip fracture: DT, drug therapy
     *hip fracture: PC, prevention
    hormone substitution
    human
    morbidity
    nausea:. . .
    DO, drug dose
     risedronic acid: DT, drug therapy
     risedronic acid: PD, pharmacology
     selective estrogen receptor modulator: DT, drug therapy
    selective estrogen receptor modulator: PD, pharmacology
       strontium ranelate: DV, drug development
     tamoxifen: DT, drug therapy
     tamoxifen: PD. pharmacology
     thiazide diuretic agent
     tibolone: DT, drug therapy
     tibolone: PD, pharmacology
     tiludronic acid: DT, drug therapy
```

tiludronic.

RN. . acid) 40391-99-9, 57248-88-1; (parathyroid hormone) 12584-96-2, 68893-82-3, 9002-64-6; (parathyroid hormone[1-34]) 12583-68-5, 52232-67-4; (raloxifene) 82640-04-8, 84449-90-1; (risedronic acid) 105462-24-6, 122458-82-6; (strontium ranelate) 135459-87-9; (tamoxifen) 10540-29-1; (tibolone) 5630-53-5; (tiludronic acid)

96538-83-9; (vitamin K group) 12001-79-5; (zoledronic acid) 118072-93-8, 131654-46-1, 165800-06-6, 165800-07-7

L9 ANSWER 9 OF 13 USPATFULL on STN

ACCESSION NUMBER: 2001:22352 USPATFULL

TITLE: Methods to improve neural outcome

INVENTOR(S): Gluckman, Peter D., Auckland, New Zealand

Williams, Christopher E., Auckland, New Zealand

Guan, Jian, Auckland, New Zealand

Aukland Uniservices Limited, Aukland, New Zealand PATENT ASSIGNEE(S):

(non-U.S. corporation)

NUMBER KIND DATE \_\_\_\_\_

PATENT INFORMATION: US 6187906 B1 20010213 APPLICATION INFO:: US 1999-332868 19990615 (9)

RELATED APPLN. INFO .: Continuation-in-part of Ser. No. US 1997-907918, filed on 11 Aug 1997

NUMBER DATE

PRIORITY INFORMATION: NZ 1998-330684 19980615 DOCUMENT TYPE: Utility

FILE SEGMENT: Granted

PRIMARY EXAMINER:

Low, Christopher S. F.

LEGAL REPRESENTATIVE: Nixon & Vanderhye

NUMBER OF CLAIMS: 11

EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 18 Drawing Figure(s); 9 Drawing Page(s)
LINE COUNT: 1057

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

. insults associated with near-miss drowning, near-miss cot death, carbon monoxide inhalation, ammonia or other gaseous intoxication, cardiac arrest, collapse, coma, meningitis, hypoglycaemia and status epilepticus; episodes of cerebral asphyxia associated with coronary bypass surgery; cerebral anoxia or ischemia associated with stroke,.

DETD . . O.sub.2 for 20 minutes, washed with 0.1M PBS (3+5 minutes) and incubated with rabbit polyclonal antisera raised against tyrosine hydroxylase (Protos Biotech, USA) diluted 1:500 with 1% goat serum for 48 hours at 4° C. The sections were washed in PBS. . .

L9 ANSWER 10 OF 13 SCISEARCH COPYRIGHT (c) 2009 The Thomson Corporation on

2001:388795 SCISEARCH ACCESSION NUMBER: THE GENUINE ARTICLE: 429BP

TITLE: Incorporation and distribution of strontium in bone

AUTHOR . Dahl S G (Reprint)

CORPORATE SOURCE: Univ Tromso, Fac Med, Dept Pharmacol, N-9037 Tromso,

Norway (Reprint) AUTHOR:

Allain P; Marie P J; Mauras Y; Boivin G; Ammann P;

Tsouderos Y; Delmas P D; Christiansen C

PUBLISHER:

CORPORATE SOURCE: CHU Angers, Lab Pharmacol & Toxicol, Angers, France; CNRS,

Lariboisiere Hosp, INSERM, U349, Paris, France; Fac Med R Laennec, INSERM, U403, Lyon, France; Univ Geneva, Hop Cantonal, Div Malad Osseuses, Dept Med Interne, CH-1211 Geneva, Switzerland; Inst Rech Int Servier, F-92415 Courbevoie, France; Ctr Clin & Basic Res, Ballerup,

Denmark

COUNTRY OF AUTHOR: Norway: France: Switzerland: Denmark SOURCE:

BONE, (APR 2001) Vol. 28, No. 4, pp. 446-453.

ISSN: 8756-3282. ELSEVIER SCIENCE INC, 655 AVENUE OF THE AMERICAS, NEW

YORK, NY 10010 USA. DOCUMENT TYPE: Article: Journal

LANGUAGE: English

REFERENCE COUNT:

Entered STN: 25 May 2001 ENTRY DATE:

Last Updated on STN: 25 May 2001

\*ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS\*

so BONE, (APR 2001) Vol. 28, No. 4, pp. 446-453.

ISSN: 8756-3282.

. . into bone has been examined in rats, monkeys, and humans after oral administration of strontium (either strontium chloride or strontium ranelate). After repeated administration for a sufficient period of time (at least 4 weeks in rats), strontium incorporation into bone reaches.

STP KeyWords Plus (R): POSTMENOPAUSAL OSTEOPOROSIS: CALCIUM-METABOLISM: MINERAL DENSITY: ILIAC BONE; RATS; FLUORIDE; RESORPTION; ARTHRITIS ; TURNOVER; SKELETON

ANSWER 11 OF 13 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on L9

ACCESSION NUMBER: 2001:550064 BIOSIS DOCUMENT NUMBER: PREV200100550064

TITLE: Strontium ranelate increases cartilage matrix

formation.

Henrotin, Y. [Reprint author]; Labasse, A. [Reprint AUTHOR(S): author]; Galais, Ph.; Tsouderos, Y.; Crielaard, J. M.

[Reprint author]; Reginster, J. Y. [Reprint author] Bone and Cartilage Metabolism Research Unit, University CORPORATE SOURCE:

Hospital, CHU Sart-Tilman, 4000, Liege, Belgium SOURCE: Clinical Rheumatology, (2001) Vol. 20, No. 5, pp.

416, print.

Meeting Info.: 5th Belgian Congress on Rheumatology.

Hasselt, Belgium. September 27-29, 2001.

CODEN: CLRHD6. ISSN: 0770-3198.

DOCUMENT TYPE: Conference; (Meeting)

Conference; Abstract; (Meeting Abstract)

LANGUAGE: English

ENTRY DATE: Entered STN: 21 Nov 2001

Last Updated on STN: 25 Feb 2002

Strontium ranelate increases cartilage matrix formation.

Clinical Rheumatology, (2001) Vol. 20, No. 5, pp. 416. print. Meeting Info.: 5th Belgian Congress on Rheumatology. Hasselt, Belgium. September 27-29, 2001.

CODEN:. . .

(Movement and Support)

Parts, Structures, & Systems of Organisms

cartilage: skeletal system, matrix formation; chondrocytes: skeletal system

SOURCE:

AB

IT Diseases

osteoarthritis: joint disease

Osteoarthritis (MeSH)

IT Chemicals & Biochemicals

insulin-like growth factor-I; interleukin-l beta; proteoglycans: production; stromelysin: activation; strontium ranelate: antiarthritic-drug

L9 ANSWER 12 OF 13 BIOSIS COPYRIGHT (c) 2009 The Thomson Corporation on STN DUPLICATE 1

ACCESSION NUMBER: 2001:120190 BIOSIS

DOCUMENT NUMBER: PREV200100120190

TITLE: Strontium ranelate increases cartilage matrix

formation.

AUTHOR(S): Henrotin, Y. [Reprint author]; Labasse, A.; Zheng, S. X.; Galais, Ph.; Tsouderos, Y.; Crielaard, J. M.; Reginster, J.

Y.

CORPORATE SOURCE: Bone and Cartilage Metabolism Research Unit Institute of

Pathology, C.H.U. Sart-Tilman, Bat B23, B-4000, Liege,

Belgium

Journal of Bone and Mineral Research, (February,

2001) Vol. 16, No. 2, pp. 299-308. print.

CODEN: JBMREJ. ISSN: 0884-0431.

DOCUMENT TYPE: Article LANGUAGE: English

ENTRY DATE: Entered STN: 7 Mar 2001 Last Updated on STN: 15 Feb 2002

TI Strontium ranelate increases cartilage matrix formation.

SO Journal of Bone and Mineral Research, (February, 2001) Vol. 16,

No. 2, pp. 299-308. print.

CODEN: JEMREJ. ISSN: 0884-0431.

Based on previous studies showing that strontium ranelate

(S12911) modulates bone loss in osteoporosis, it could be hypothesized that this drug also is effective on cartilage degradation in osteoarthritis (OR). This was investigated in vitro on normal and OA human chondrocytes treated or not treated with interleukin-lbeta (IL-lbeta). This. . in OA cartilage. Chondrocytes were isolated from cartilage by enzymatic digestion and cultured for 24-72 h with 10-4-10-3 M strontium reneltet, 10-3 M calcium reneltete, 10-3 M calcium reneltete. (or 2.10-3 M SrC12 with or without IL-lbeta or insulin-like growth factor I (IGF-I). Stromelysin activity and stromelysin quantitation were. .

were quantified by labeled sulfate (Na235504) incorporation. This method allowed the PC size after exclusion chromatography to be determined. Strontium ranelate, calcium ranelate, and Src12 did

not modify stromelysin synthesis even in the presence of IL-lbeta. Calcium ranelate induced stromelysin activation whereas

strontium compounds were ineffective. Strontium <u>ranelate</u> and SrC12 both strongly stimulated PG production suggesting an ionic effect of strontium independent of the organic moiety. Moreover, 10-3 M strontium <u>ranelate</u> increased the stimulatory effect of IGF-1 (10-9 M) on PG

synthesis but did not reverse the inhibitory effect of IL-lbeta. Strontium <u>ranelate</u> strongly stimulates human cartilage matrix formation in vitro by a direct ionic effect without stimulating the chondroresorption processes. This finding provides a preclinical basis

for in vivo testing of strontium ranelate in OA.

IT Parts, Structures, & Systems of Organisms

cartilage: skeletal system; chondrocytes: skeletal system

IT Diseases

## osteoarthritis: joint disease Osteoarthritis (MeSH)

IT Diseases

osteoporosis: bone disease Osteoporosis (MeSH)

IT Chemicals & Biochemicals

interleukin-1-beta; strontium ranelate [S12911]

RN 135459-87-9 (S12911)

L9 ANSWER 13 OF 13 IMSPRODUCT COPYRIGHT 2009 IMSWORLD on STN

SO Drug Launches, (<u>20</u> <u>Sep</u> <u>1999</u>)

CN Trade Name: PROTOS
CN Chemical Name: protoporphyrin IX disodium

TX Hepatic disorders caused cholecystitis or gall stones